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HASKELL LABORATORY DISCOVERY TOXICOLOGY GROUP

In Vitro Rat Hepatocyte Screen

WR:	17199
SERVICE CODE:	1599
HASKELL#:	28072
STUDY COMPLETED:	12-Jun-07
NOTEBOOK #:	E-111389-AT

STUDY DESIGN:

Test Substance:	HFPO Dimer Acid Salt
Species:	Rat
Strain:	Crl:CD®(SD)IGS BR
Gender:	Male and Female
Cell Concentration:	1×10^6 cells/mL (clearance incubations) 5×10^6 cells/mL (biotransformation incubations)
Reaction Buffer:	L-15 medium
pH:	7.4
Reaction Volume:	2.5 mL
Dose Vehicle:	Nanopure Water
Dose Volume:	10 µL/mL
Final Concentration:	2 µM = 694 ppb (clearance incubations) 200 µM = 69.4 ppm (biotransformation incubations)
Replicates/Sex:	3 test, 3 heat –inactivated controls, 1 biotransformation, 1 positive control (4-nonylphenol).
Time Points:	5, 15, 30, 45, 60, 90, and 120 minutes
Incubation Temperature:	37ºC
Extraction:	1:2, Sample:Acetonitrile
Dilution:	1:1, Sample:Nanopure Water
Final Dilution Factor:	6x
Analytical:	LC/MS

OBJECTIVE:

To estimate metabolic clearance of test compound in rat hepatocytes and extrapolate results to whole animal and to identify metabolites and describe probable metabolic pathways for the compound tested.

PARAMETERS:

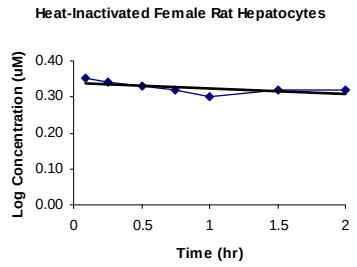
Half-life (T1/2), in vitro clearance (mL/hr), extrapolated in vivo clearance (mL/hr/kg), and metabolite ID.

RESULTS:

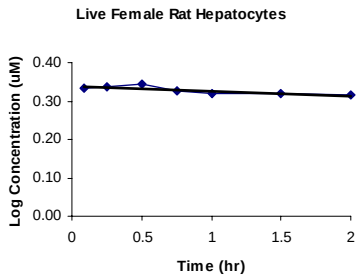
Summary: No apparent loss of the parent compound was observed within 2 hours of incubation compared to heat-inactivated controls. No metabolites were identified in the biotransformation incubation samples.

Representative Figures

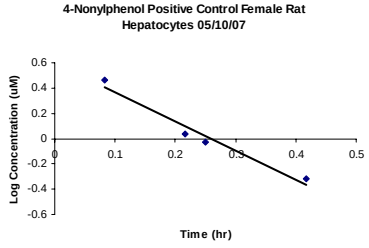
Heat-Inactivated Female Control



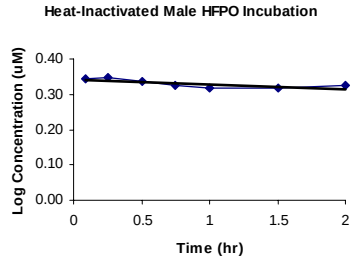
Test Substance Female



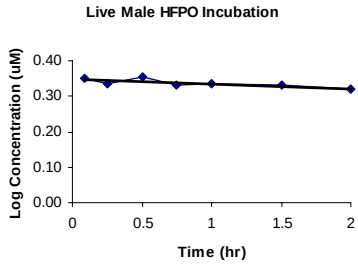
Positive Control Female



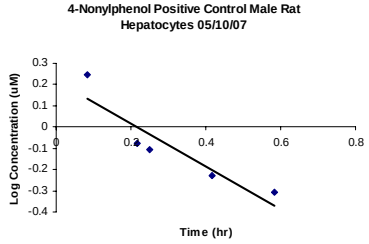
Heat-Inactivated Male Control



Test Substance Male



Positive Control Male



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